

SEARCH REQUEST FORM

Scientific and Technical Information Center

Requester's Full Name: K. Weddington Examiner #: 68082 Date: 4-28-03
 Art Unit: 1114 Phone Number 308-4650 Serial Number: 09/926,807
 Mail Box and Bldg/Room Location: SM1-2A17 Results Format Preferred (circle): PAPER DISK E-MAIL

If more than one search is submitted, please prioritize searches in order of need. *ME*

Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc., if known. Please attach a copy of the cover sheet, pertinent claims, and abstract.

Title of Invention: _____

Inventors (please provide full names): Jacobus Johnannes Wannie Meyer, Namrita Lall

Earliest Priority Filing Date: _____

For Sequence Searches Only Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

A method of treating tuberculosis with formula I

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Type of Search	Vendors and cost where applicable
NA Sequence (#)	STN _____
AA Sequence (#)	Dialog _____
Structure (#)	Questel/Orbit _____
Bibliographic	Dr.Link _____
Litigation	Lexis/Nexis _____
Fulltext	Sequence Systems _____
Patent Family	WWW/Internet _____
Other	Other (specify) _____

Searcher: Sheppard

Searcher Phone #: 308-4499

Searcher Location: _____

Date Searcher Picked Up: _____

Date Completed: 4/30/03

Searcher Prep & Review Time: _____

Clerical Prep Time: _____

Online Time: _____

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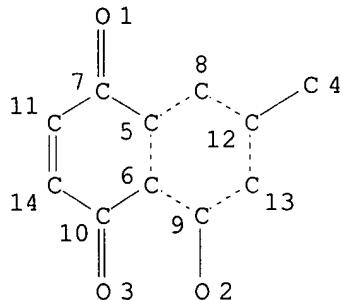
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FILE COVERS 1907 - 30 Apr 2003 VOL 138 ISS 18
FILE LAST UPDATED: 29 Apr 2003 (20030429/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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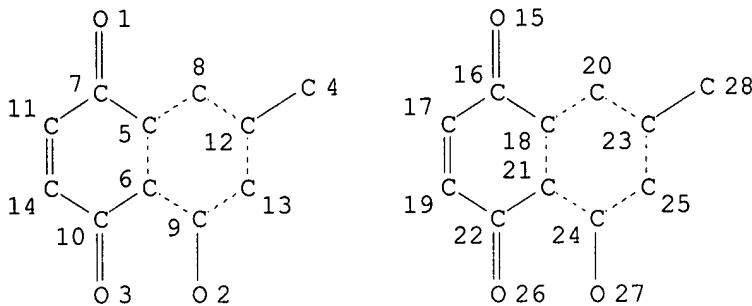
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L24 STR



NODE ATTRIBUTES:
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DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
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NUMBER OF NODES IS 14

STEREO ATTRIBUTES: NONE
L25 875 SEA FILE=REGISTRY SSS FUL L24
L26 STR



NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 28

STEREO ATTRIBUTES: NONE

L27 112 SEA FILE=REGISTRY SUB=L25 SSS FUL L26

L28 160 SEA FILE=HCAPLUS ABB=ON PLU=ON L27

L29 2 SEA FILE=HCAPLUS ABB=ON PLU=ON L28 AND ?TUBERCUL?

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=> d ibib abs hitrn 129 1-2

L29 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:794683 HCAPLUS

DOCUMENT NUMBER: 137:75724

TITLE: Inhibition of drug-sensitive and drug-resistant strains of *Mycobacterium tuberculosis* by diospyrin, isolated from *Euclea natalensis*

AUTHOR(S): Lall, N.; Meyer, J. J. M.

CORPORATE SOURCE: Department of Botany, University of Pretoria, Pretoria, 0002, S. Afr.

SOURCE: Journal of Ethnopharmacology (2001), 78(2-3), 213-216
CODEN: JOETD7; ISSN: 0378-8741

PUBLISHER: Elsevier Science Ireland Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The binaphthoquinoid, diospyrin, was isolated from *Euclea natalensis* A.D.C., and evaluated for its activity against drug-sensitive and drug-resistant strains of *Mycobacterium tuberculosis*. The minimal inhibitory concn. (MIC) of diospyrin was found to be 100 .mu.g/mL for all the *M. tuberculosis* strains.

IT 28164-57-0P, Diospyrin

RL: PAC (Pharmacological activity); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(M. *tuberculosis* drug-sensitive and drug-resistant strains inhibition by *Euclea natalensis* isolate diospyrin)

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:12397 HCAPLUS

DOCUMENT NUMBER: 134:68700
 TITLE: Naphthoquinone derivatives and their use in the treatment and control of **tuberculosis**
 INVENTOR(S): Meyer, Jacobus Johannes Marion; Lall, Namrita
 PATENT ASSIGNEE(S): University of Pretoria, S. Afr.
 SOURCE: PCT Int. Appl., 22 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001000554	A2	20010104	WO 2000-IB837	20000622
WO 2001000554	A3	20010705		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1194137	A2	20020410	EP 2000-937123	20000622
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
PRIORITY APPLN. INFO.:			ZA 1999-4176	A 19990624
			WO 2000-IB837	W 20000622

OTHER SOURCE(S): MARPAT 134:68700
 AB Naphthoquinone derivs., or pharmaceutically acceptable salts thereof, are useful for the treatment and/or control of a **tuberculosis** caused by *Mycobacterium tuberculosis* in a patient. The naphthoquinone derivs. are administered orally, i.v., i.m., or transdermally. For example, diospyrin and 7-methyljuglone controlled effectively the sensitive and resistant strains of *M. tuberculosis* with min. inhibitory concn. (MIC) of 0.1 .mu.g/mL for diospyrin and 50 .mu.g/mL for 7-methyljuglone, while the MIC for the combination of two drugs was 10 .mu.g/mL.
 IT 28164-57-0, Diospyrin
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (naphthoquinone derivs. and their use in control and treatment of **tuberculosis**)

=> select hit rn 129 1-2
 E25 THROUGH E25 ASSIGNED

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 FILE 'REGISTRY' ENTERED AT 16:46:17 ON 30 APR 2003
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STRUCTURE FILE UPDATES: 29 APR 2003 HIGHEST RN 507441-92-1
 DICTIONARY FILE UPDATES: 29 APR 2003 HIGHEST RN 507441-92-1

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2003

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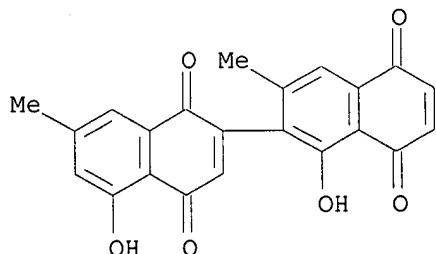
Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

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=> s e25
L30 1 28164-57-0/BI
(28164-57-0/RN)

=> d ide can 130 1

L30 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
RN 28164-57-0 REGISTRY
CN [2,2'-Binaphthalene]-1,4,5',8'-tetrone, 1',5-dihydroxy-3',7-dimethyl-
(8CI, 9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
CN Diopspyrin (7CI)
OTHER NAMES:
CN Euclein
FS 3D CONCORD
DR 27939-56-6
MF C22 H14 O6
LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, DDFU, DRUGU, EMBASE,
IPA, MEDLINE, NAPRALERT, RTECS*, SPECINFO, TOXCENTER
(*File contains numerically searchable property data)



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

54 REFERENCES IN FILE CA (1957 TO DATE)
2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
54 REFERENCES IN FILE CAPLUS (1957 TO DATE)
2 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 138:251352

REFERENCE 2: 137:140377

REFERENCE 3: 137:75724

REFERENCE 4: 134:68700
REFERENCE 5: 133:168474
REFERENCE 6: 133:28470
REFERENCE 7: 133:17317
REFERENCE 8: 132:305640
REFERENCE 9: 132:134818
REFERENCE 10: 130:164730

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FILE LAST UPDATED: 29 Apr 2003 (20030429/ED)

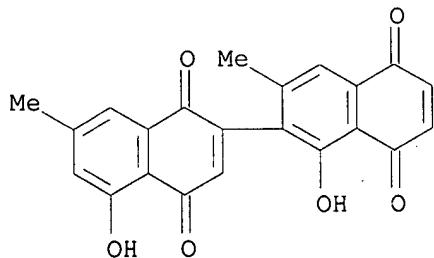
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L28 160 SEA FILE=HCAPLUS ABB=ON PLU=ON L27
L29 2 SEA FILE=HCAPLUS ABB=ON PLU=ON L28 AND ?TUBERCUL?
L31 4 SEA FILE=HCAPLUS ABB=ON PLU=ON L28(L) (?MEDIC? OR ?PHARM? OR
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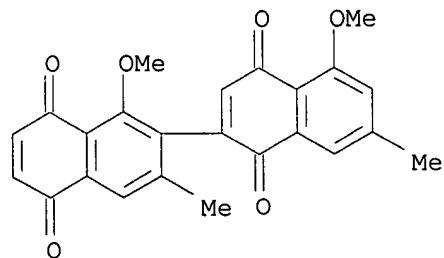
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L32 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2003 ACS
ACCESSION NUMBER: 2000:145304 HCAPLUS
DOCUMENT NUMBER: 132:305640
TITLE: Effects of atovaquone and diospyrin-based drugs on the
cellular ATP of *Pneumocystis carinii* f. sp. *carinii*
AUTHOR(S): Cushion, Melanie T.; Collins, Margaret; Hazra,
Banasri; Kaneshiro, Edna S.
CORPORATE SOURCE: Department of Internal Medicine, University of
Cincinnati College of Medicine, and Veterans Affairs
Medical Center, Cincinnati, OH, 45267-0560, USA
SOURCE: *Antimicrobial Agents and Chemotherapy* (2000), 44(3),
713-719
CODEN: AMACCQ; ISSN: 0066-4804
PUBLISHER: American Society for Microbiology
DOCUMENT TYPE: Journal
LANGUAGE: English
AB Atovaquone (also called Mepron, or 566C80) is a naphthoquinone used for
the treatment of infections caused by pathogens such as *Plasmodium* spp.

and *Pneumocystis carinii*. The mechanism of action against the malarial parasite is the inhibition of dihydroorotate dehydrogenase (DHOD), a consequence of blocking electron transport by the drug. As an analog of ubiquinone (coenzyme Q [CoQ]), atovaquone irreversibly binds to the mitochondrial cytochrome bcl complex; thus, electrons are not able to pass from dehydrogenase enzymes via CoQ to cytochrome c. Since DHOD is a crit. enzyme in pyrimidine biosynthesis, and because the parasite cannot scavenge host pyrimidines, the drug is lethal to the organism. Oxygen consumption in *P. carinii* is inhibited by the drug; thus, electron transport has also been identified as the drug target in *P. carinii*. However, unlike *Plasmodium* DHOD, *P. carinii* DHOD is inhibited only at high atovaquone concns., suggesting that the organism may salvage host pyrimidines and that atovaquone exerts its primary effects on ATP biosynthesis. In the present study, the effect of atovaquone on ATP levels in *P. carinii* was measured directly from 1 to 6 h and then after 24, 48, and 72 h of exposure. The av. 50% inhibitory concn. after 24 to 72 h of exposure was 1.5 .mu.g/mL (4.2 .mu.M). The kinetics of ATP depletion were in contrast to those of another family of naphthoquinone compds., diospyrin and two of its derivs. Whereas atovaquone reduced ATP levels within 1 h of exposure, the diospyrins required at least 48 h. After 72 h, the diospyrins were able to decrease ATP levels of *P. carinii* at nanomolar concns. These data indicate that although naphthoquinones inhibit the electron transport chain, the mol. targets in a given organism are likely to be distinct among members of this class of compds.

IT 28164-57-0, Diospyrin 39093-14-6, Diospyrin dimethylether
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)
 (effects of atovaquone and diospyrin-based drugs on cellular ATP of *Pneumocystis carinii*)
 RN 28164-57-0 HCPLUS
 CN [2,2'-Binaphthalene]-1,4,5',8'-tetrone, 1',5-dihydroxy-3',7-dimethyl-
 (8CI, 9CI) (CA INDEX NAME)



RN 39093-14-6 HCPLUS
 CN [2,2'-Binaphthalene]-1,4,5',8'-tetrone, 1',5-dimethoxy-3',7-dimethyl-
 (9CI) (CA INDEX NAME)



REFERENCE COUNT:

37

THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L32 ANSWER 2 OF 3 HCPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 1998:27367 HCPLUS
 DOCUMENT NUMBER: 128:162607
 TITLE: Cell line-directed screening assay for inhibitors of thioredoxin reductase signaling as potential anti-cancer drugs
 AUTHOR(S): Kunkel, Mark W.; Kirkpatrick, D. Lynn; Johnson, Jill I.; Powis, Garth
 CORPORATE SOURCE: Arizona Cancer Center, University of Arizona Health Sciences Center, Tucson, AZ, 85724-5024, USA
 SOURCE: Anti-Cancer Drug Design (1997), 12(8), 659-670
 CODEN: ACDDEA; ISSN: 0266-9536
 PUBLISHER: Oxford University Press
 DOCUMENT TYPE: Journal
 LANGUAGE: English

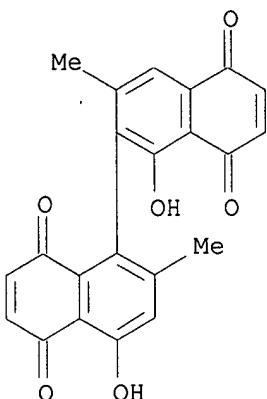
AB We have used a cell line-directed screening approach (CDSA) to identify novel inhibitors of the thioredoxin reductase signaling pathway which contributes to the transformed phenotype of some human tumors. Two 2-imidazolyl disulfide compds., previously identified as inhibitors of thioredoxin reductase, were screened for growth inhibitory activity in the National Cancer Institute (NCI) human cancer cell line panel. The COMPARE pattern recognition algorithm was used to identify similar compds. from >60,000 compds. in the NCI investigational drug database. Of 47 nondiscreet compds. tested in a thioredoxin reductase/thioredoxin insulin redn. assay, 37 (77%) were inhibitors with IC50s < 10 μ g/mL and 15 of those (32%) had IC50s < 1 μ g/mL. These compds. were all as selective or more selective for thioredoxin reductase than for glutathione reductase, while three compds. were inhibitors of thioredoxin. In comparison to CDSA, the no. of compds. with IC50s < 1 μ g/mL identified by screening of 52 compds. from the database whose growth inhibiting activity was unrelated to the activity of the disulfide compds. was only 2%. Screening of 221 randomly selected natural products gave only 3% of compds. with IC50s < 1 μ g/mL. Thus, the CDSA using data from the NCI cancer cell panel and known inhibitors of the selected target as seed compds. can greatly increase hit rates, compared with random screening, for identifying novel inhibitors of a target, in this case thioredoxin signaling.

IT 89475-33-2
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(cell line-directed screening assay for inhibitors of thioredoxin reductase signaling as potential anti-cancer drugs)

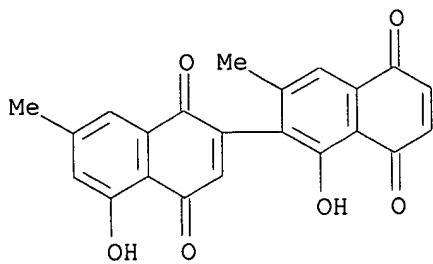
RN 89475-33-2 HCPLUS

CN [1,2'-Binaphthalene]-5,5',8,8'-tetrone, 1',4-dihydroxy-2,3'-dimethyl- (9CI) (CA INDEX NAME)

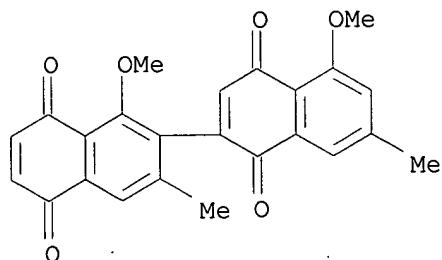


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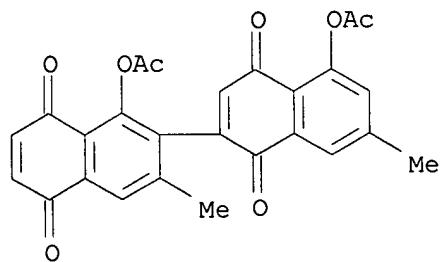
L32 ANSWER 3 OF 3 HCPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 1996:542516 HCPLUS
 DOCUMENT NUMBER: 125:237771
 TITLE: Pharmacological studies on the effect of the treatment of Swiss A mice with diospyrin, a tumor-inhibitory plant product, and its synthetic derivatives
 AUTHOR(S): Pal, Sampa; Banerjee, Amalendu; Hazra, Banasri; Ray, Ratnamala; Bhattacharya, Dilip K.
 CORPORATE SOURCE: Dep. Pharmacy Chem., Jadavpur Univ., Calcutta, 700 032, India
 SOURCE: Phytotherapy Research (1996), 10(5), 393-397
 CODEN: PHYREH; ISSN: 0951-418X
 PUBLISHER: Wiley
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Diospyrin, a bisnaphthoquinonoid plant product, and its derivs., have shown significant inhibitory activities against murine tumors in vivo. Studies on the hematol. status, serum protein and creatinine levels, activities of several serum glycolytic enzymes, and histopathol. of the mice inoculated with Ehrlich ascites carcinoma were carried out after treatment with diospyrin and four synthetic derivs. The prognostic significance of the pharmacol. parameters acting as markers of the diseased state was evident from these findings. Normal mice were also studied before and after treatment with these compds. which did not cause noticeable adverse effects on the vital parameters, thereby indicating the possibility of the utilization of diospyrin and derivs. as appropriate therapeutic agents.
 IT 28164-57-0, Diospyrin 39093-14-6
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (pharmacol. studies on effect of treatment of Swiss A mice with antitumor agent diospyrin and synthetic derivs.)
 RN 28164-57-0 HCPLUS
 CN [2,2'-Binaphthalene]-1,4,5',8'-tetrone, 1',5-dihydroxy-3',7-dimethyl- (8CI, 9CI) (CA INDEX NAME)



RN 39093-14-6 HCPLUS
 CN [2,2'-Binaphthalene]-1,4,5',8'-tetrone, 1',5-dimethoxy-3',7-dimethyl-
 (9CI) (CA INDEX NAME)



IT 60544-03-8
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (pharmacol. studies on effect of treatment of normal and tumor-bearing mice with antitumor agent diospyrin and synthetic derivs.)
 RN 60544-03-8 HCPLUS
 CN [2,2'-Binaphthalene]-1,4,5',8'-tetrone, 1',5-bis(acetyloxy)-3',7-dimethyl-
 (9CI) (CA INDEX NAME)



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L33      3 SEA FILE=HCAPLUS ABB=ON PLU=ON L31 NOT L29
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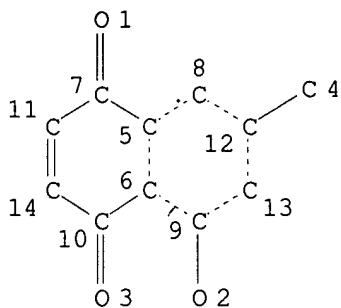
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L36 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2003 ACS
ACCESSION NUMBER: 2003:89327 HCAPLUS
TITLE: Antimycobacterial activity of **diospyrin**
derivatives and a structural analogue of
diospyrin against **Mycobacterium**
tuberculosis *in vitro*
AUTHOR(S): Lall, N.; Das Sarma, M.; Hazra, B.; Meyer, J. J. M.
CORPORATE SOURCE: Department of Botany, University of Pretoria,
Pretoria, 0002, S. Afr.
SOURCE: Journal of Antimicrobial Chemotherapy (2003), 51(2),
435-438
CODEN: JACHDX; ISSN: 0305-7453
PUBLISHER: Oxford University Press
DOCUMENT TYPE: Journal
LANGUAGE: English
AB Three derivs. and one structural analog of **diospyrin** were
synthesized and investigated for their inhibitory activity against
Mycobacterium **tuberculosis** employing the rapid radiometric
method *in vitro*. A novel aminoacetate deriv. was found to be more active
than the parent compd., the MICs being 50 and 100 mg/L, resp., for a
drug-susceptible strain, H37Rv, of **M. tuberculosis**. This deriv.
also exhibited an MIC of 50 mg/L for a few multidrug-resistant strains of
M. tuberculosis. The other two derivs. and the analog did not
show any significant antimycobacterial activity at the highest concn. (100
mg/L) tested.
REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L24 STR



NODE ATTRIBUTES:

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DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

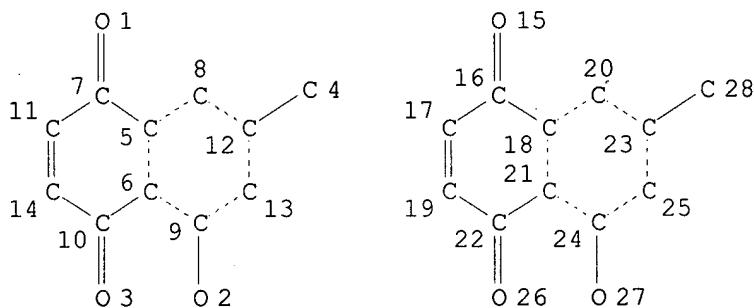
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NUMBER OF NODES IS 14

STEREO ATTRIBUTES: NONE

L25 875 SEA FILE=REGISTRY SSS FUL L24

L26 STR



NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

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NUMBER OF NODES IS 28

STEREO ATTRIBUTES: NONE

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L31 4 SEA FILE=HCAPLUS ABB=ON PLU=ON L28(L) (?MEDIC? OR ?PHARM? OR ?DRUG? OR THERAP? OR MYCOBACTERIUM)

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L34 68 SEA FILE=HCAPLUS ABB=ON PLU=ON L33

L35 3 SEA FILE=HCAPLUS ABB=ON PLU=ON L34 AND ?TUBERCU?

L36 1 SEA FILE=HCAPLUS ABB=ON PLU=ON L35 NOT (L29 OR L32)

L40 5 SEA FILE=HCAPLUS ABB=ON PLU=ON L28 AND (?MYCOBACT? OR ANTIBACT?)

L41 3 SEA FILE=HCAPLUS ABB=ON PLU=ON L40 NOT (L29 OR L32 OR L36)

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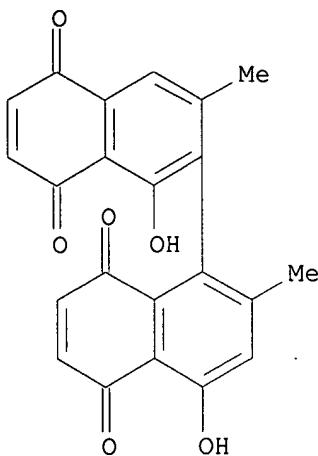
=> d ibib abs hitstr 141 1-3

L41 ANSWER 1 OF 3 HCPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 2000:243113 HCPLUS
 DOCUMENT NUMBER: 133:28470
 TITLE: **Antibacterial activity of diospyrin, isodiospyrin and bisisodiospyrin from the root of Diospyros piscatoria (Gurke) (Ebenaceae)**
 AUTHOR(S): Adeniyi, B. A.; Fong, H. H. S.; Pezzuto, J. M.; Luyengi, L.; Odelola, H. A.
 CORPORATE SOURCE: Department of Pharmaceutical Microbiology and Clinical Pharmacy, College of Medicine, University of Ibadan, Ibadan, Nigeria
 SOURCE: Phytotherapy Research (2000), 14(2), 112-117
 CODEN: PHYREH; ISSN: 0951-418X
 PUBLISHER: John Wiley & Sons Ltd.
 DOCUMENT TYPE: Journal
 LANGUAGE: English

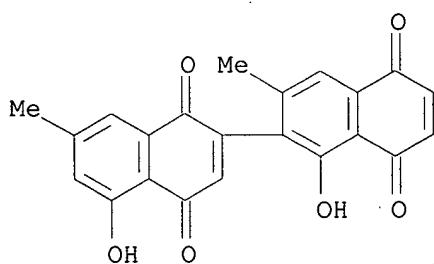
AB Two dimeric naphthoquinones, diospyrin and isodiospyrin, isolated from the root of *Diospyros piscatoria* (Gurke), a common ingredient in several folk medicines, have been shown to have a broad spectrum of **antibacterial** activity. The min. inhibitory concns. (MICs) of diospyrin against *Streptococcus pyogenes* ATCC 12344 and *Streptococcus pneumoniae* ATCC 33400 ranged from 1.56 to 50 .mu.g/mL. While those against *Salmonella choleraesuis* serotype *typhi* (*S. typhi*), ATCC 6539 and **Mycobacterium chelonae** ATCC 19977 were between 25 and 100 .mu.g/mL. Isodiospyrin was more active than its racemic isomer diospyrin. The MICs against Gram-pos. bacteria ranged from 0.78 to 50 .mu.g/mL. While those against *Pseudomonas aeruginosa* ATCC 15443 and *S. typhi* ranged from 50 to 100 .mu.g/mL. The MIC for *M. chelonae* was between 6.25 and 25 .mu.g/mL. MICs were found to increase with the concn. of cells used for the inoculum. The MICs for *Bacillus subtilis* ATCC 6633 increased up to the highest concn. of cells tested. The same phenomenon was obsd. on *M. chelonae*, but with better effect in the latter. The kinetics of bacteria studies against both *B. subtilis* and *M. chelonae* increases with increasing concn. of isodiospyrin tested. Two tetrameric forms of plumbagin were isolated. The naphthoquinone bisisodiospyrin, gave MIC values between 300 and 400 .mu.g/mL. The second, as yet unidentified tetramer, was not active at 500 .mu.g/mL.

IT 20175-84-2P, Isodiospyrin 28164-57-0P, Diospyrin
 30276-87-0P, Bisodiospyrin
 RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)
 (**antibacterial** activity of diospyrin, isodiospyrin, and bisisodiospyrin from the root of *Diospyros piscatoria*)

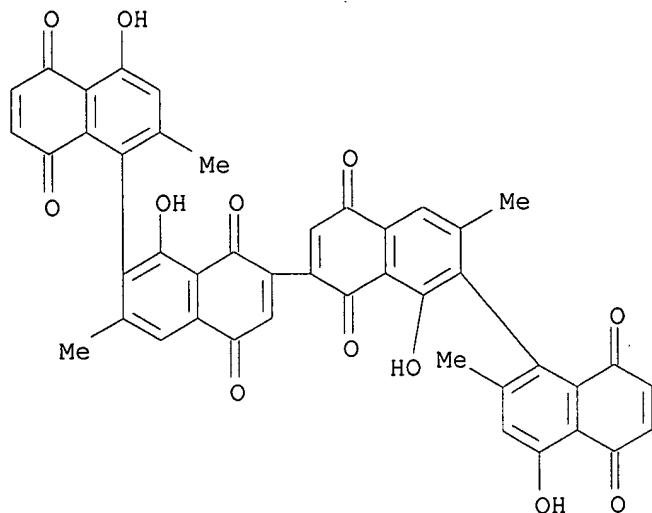
RN 20175-84-2 HCPLUS
 CN [1,2'-Binaphthalene]-5,5',8,8'-tetrone, 1',4-dihydroxy-2,3'-dimethyl-, (1R)- (9CI) (CA INDEX NAME)



RN 28164-57-0 HCPLUS
CN [2,2'-Binaphthalene]-1,4,5',8'-tetrone, 1',5-dihydroxy-3',7-dimethyl- (8CI, 9CI) (CA INDEX NAME)



RN 30276-87-0 HCPLUS
CN [1,2':7',2'':7'':1''''-Quaternaphthalene]-1'',4'',5,5',5''',8,8',8''''-octone, 1',4,4'',8''-tetrahydroxy-2,2'',3',6''-tetramethyl- (9CI) (CA INDEX NAME)

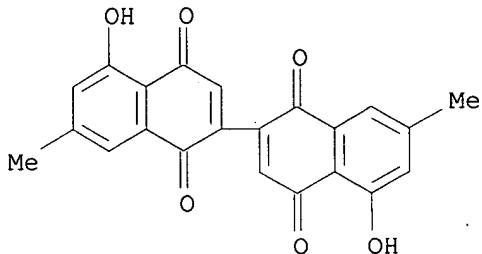


REFERENCE COUNT:

20

THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L41 ANSWER 2 OF 3 HCPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 1999:430463 HCPLUS
 DOCUMENT NUMBER: 131:291097
 TITLE: Constituents of *Diospyros lolin*, *D. Maritima* and *D. Novoguinensis*
 AUTHOR(S): Khan, M. R.; Timi, D.
 CORPORATE SOURCE: Department of Applied Sciences, Papua New Guinea
 University of Technology, Papua, Papua New Guinea
 SOURCE: *Fitoterapia* (1999), 70(2), 194-196
 CODEN: FTRPAE; ISSN: 0367-326X
 PUBLISHER: Elsevier Science B.V.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Antibacterial activity of 7-methyljuglone (I), plumbagin (II), and biramentaceone isolated from *Diospyros* species was studied. Only I and II showed antibacterial activity.
 IT 24456-79-9
 RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)
 (isolation and antibacterial activity of constituents of *Diospyros*)
 RN 24456-79-9 HCPLUS
 CN [2,2'-Binaphthalene]-1,1',4,4'-tetrone, 5,5'-dihydroxy-7,7'-dimethyl- (8CI, 9CI) (CA INDEX NAME)



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L41 ANSWER 3 OF 3 HCPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 1979:98323 HCPLUS
 DOCUMENT NUMBER: 90:98323
 TITLE: Mutagenicity and antibacterial activity of mycotoxins produced by *Penicillium islandicum* Sopp and *Penicillium rugulosum*
 AUTHOR(S): Stark, A. A.; Townsend, J. M.; Wogan, G. N.; Demain, A. L.; Manmade, A.; Ghosh, A. C.
 CORPORATE SOURCE: Dep. Nutr. Food Sci., Massachusetts Inst. Technol., Cambridge, MA, USA
 SOURCE: *Journal of Environmental Pathology and Toxicology* (1978), 2(2), 313-24
 CODEN: JEPTDQ; ISSN: 0146-4779
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Twelve mycotoxins produced by *P. islandicum* and *P. rugulosum* in solid-state fermn. on grains were purified and tested for mutagenicity and antibacterial activity in *Salmonella*/mammalian microsome assays. The mutations studied were reversions of histidine auxotrophs to prototrophy in strains TA98 and TA100 and forward mutations to 8-azaguanine resistance (8AGR) in strain TM677. Rubroskyrin [21884-47-9], (+)rugulosin [23537-16-8], lumiluteoskyrin [

22333-61-5] (a photoproduct of (-)luteoskyrin [21884-44-6]), and simatoxin [66257-36-1] (a new water-sol. metabolite of unknown structure) induced 8AGR mutations in strain TM677 but not histidine reversions in strains TA98 and TA100. Mutagenic potency was reduced by rat-liver microsomes. The carcinogens (-)luteoskyrin and cyclochlorotine [12663-46-6] were **antibacterial** but not mutagenic. (+)Rugulosin, rubroskyrin, lumiluteoskyrin, and high concns. of simotoxin were also **antibacterial**. **Antibacterial** activity but not mutagenicity was obsd. with pibasterol [66257-37-2] and skyrin [602-06-2]. Chrysophanol [481-74-3], islandicin [476-56-2], iridoskyrin [568-42-3], and emodin [518-82-1] were inactive as mutagens or as **antibacterial** agents.

IT 22333-61-5

RL: BIOL (Biological study)

(of *Penicillium islandicum* and *Penicillium rugulosum*, bactericidal action and mutagenicity of)

RN 22333-61-5 HCPLUS

CN 7,17:8,16-Dimethanocyclodeca[1,2-b:5,6-b']dinaphthalene-5,6,9,10,15,18-hexone, 7,8,16,17-tetrahydro-1,4,11,14,19,20-hexahydroxy-2,13-dimethyl- (8CI, 9CI) (CA INDEX NAME)

